

FORM PTO-1449 FIRST SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT	ATTY. DOCKET NO. 2072.0010002/LBB/SJE	APPLICATION NO. To Be Assigned (Div. of 09/984,664; Filed: October 30, 2001)
	APPLICANT HANNA, Michelle M.	
	FILING DATE Herewith (June 24, 2003)	GROUP To Be Assigned

U.S. PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE
	AA1	3,996,345	12/1976	Ullman et al.			
	AB1	4,351,760	09/1982	Khanna et al.			
	AC1	4,582,788	04/1986	Erlich			
	AD1	4,683,194	07/1987	Saiki et al.			
	AE1	4,739,044	04/1988	Stabinsky			
	AF1	4,757,141	07/1988	Fung et al.			
	AG1	4,786,600	11/1988	Kramer et al.			
	AH1	5,130,238	07/1992	Malek et al.			
	AI1	5,169,766	12/1992	Schuster et al.			
	AJ1	5,508,178	04/1996	Rose et al.			
	AK1	5,595,891	01/1997	Rose et al.			

FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB- CLASS	TRANSLATION
	AL1						Yes No
	AM1						Yes No
	AN1						Yes No
	AO1						Yes No
	AP1						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

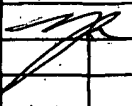

	AR	1	Agrawal, S.; and Tang, J.-Y., "Site-specific functionalization of oligodeoxynucleotides for non-radioactive labeling," Tetrahedron Lett. 31:1543-1546, Pergamon Press (1990)
	AS	1	Chamberlin, M.J., "Bacterial DNA-Dependent RNA Polymerases," in The Enzymes, Boyer, P.D., ed., Academic Press, New York, N.Y. pp. 61, 84-86 (1982)
	AT	1	Costas, C., et al., "RNA-protein crosslinking to AMP residues at internal positions in RNA with a new photocrosslinking ATP analog," Nucleic Acids Res. 28:1849-1858, Oxford University Press (May 2000)

EXAMINER	DATE CONSIDERED 2-16-06
EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to Applicant.	

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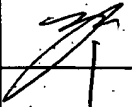

U.S. PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE
	AA2	5,648,211	07/1997	Fraiser et al.			
	AB2	5,654,142	08/1997	Kievits et al.			
	AC2	5,679,512	10/1997	Laney et al.			
	AD2	5,683,879	11/1997	Laney et al.			
	AE2	5,744,308	04/1998	Guillou-Bonnici et al.			
	AF2	5,766,849	06/1998	McDonough et al.			
	AG2	5,824,517	10/1998	Cleuziat et al.			
	AH2	5,858,801	01/1999	Brizzolara			
	AI2	6,008,334	12/1999	Hanna			
	AJ2	6,107,037	08/2000	Sousa et al.			
	AK2	6,107,039	08/2000	Hanna			

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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	2	Daube, S.S., and von Hippel, P.H., "Functional Transcription Elongation Complexes from Synthetic RNA-DNA Bubble Duplexes," Science 258:1320-1324, American Association for the Advancement of Science (1992)				
	AS	2	Dunn, J.J., and Studier, F.W., "Complete Nucleotide Sequence of Bacteriophage T7 DNA and the Locations of T7 Genetic Elements," J. Mol. Biol. 166:477-535, Academic Press, Inc. (1983)				
	AT	2	Gait, M.J., et al., "Oligoribonucleotide synthesis," in Oligonucleotides and Analogues, Eckstein, F., ed., Oxford University Press, Oxford, Great Britain, pp. 25-31 (1992)				

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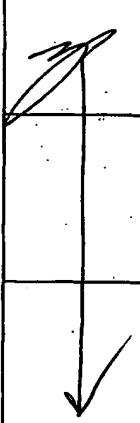
	AR	3	Gait, M.J., "An Introduction to Modern Methods of DNA Synthesis," in <i>Oligonucleotide synthesis: a practical approach</i> , Gait, M.J., ed., Oxford University Press, Oxford, Great Britain, pp.1-22 (1984)
	AS	3	Geider, K., et al., "An RNA transcribed from DNA at the origin of phage fd single strand to replicative form conversion," <i>Proc. Natl. Acad. Sci. USA</i> 75:645-649, National Academy Press (1978)
	AT	3	Guatelli, J.C., et al., "Isothermal, in vitro amplification of nucleic acids by a multienzyme reaction modeled after retroviral replication," <i>Proc. Natl. Acad. Sci. USA</i> 87:1874-1878, National Academy Press (1990)

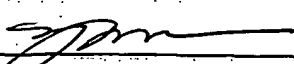
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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)							
	AR	4	Gupta, K.C., et al., "A general method for the synthesis of 3'-sulfhydryl and phosphate group containing oligonucleotides," <i>Nucleic Acids Res.</i> 19:3019-3025, Oxford University Press (1991).				
	AS	4	Gurevich, V.V., et al., "Preparative in Vitro mRNA Synthesis Using SP6 and T7 RNA Polymerases," <i>Anal. Biochem.</i> 195:207-213, Academic Press, Inc. (1991)				
	AT	4	Hanna, M.M., "Photoaffinity Cross-Linking Methods for Studying RNA-Protein Interactions," <i>Methods Enzymol.</i> 180:383-409, Academic Press, Inc. (1989)				

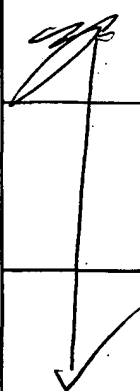
EXAMINER 	DATE CONSIDERED 7-16-06
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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	5	Hanna, M.M., et al., "Probing the environment of nascent RNA in <i>Escherichia coli</i> transcription elongation complexes utilizing a new fluorescent ribonucleotide analog," <i>Nucleic Acids Res.</i> 27:1369-1376, Oxford University Press (1999)				
	AS	5	Hanna, M.M., et al., "Synthesis and characterization of a new photocrosslinking CTP analog and its use in photoaffinity labeling <i>E.coli</i> and T7 RNA polymerases," <i>Nucleic Acids Res.</i> 21:2073-2079, Oxford University Press (1993)				
	AT	5	He, B., et al., "Preparation of probe-modified RNA with 5-mercapto-UTP for analysis of protein-RNA interactions," <i>Nucleic Acids Res.</i> 23:1231-1238, Oxford University Press (1995)				

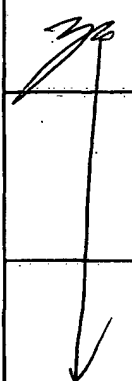
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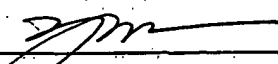
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	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)							
	AR	6	Kwoh, D.Y., et al., "Transcription-based amplification system and detection of amplified human immunodeficiency virus type 1 with a bead-based sandwich hybridization format," Proc. Natl. Acad. Sci. USA 86:1173-1177, National Academy Press (1989)				
	AS	6	Lewis, M.K., and Burgess, R.R., "Transcription of Simian Virus 40 DNA by Wheat Germ RNA Polymerase II," J. Biol. Chem. 255:4928-4936, American Society for Biochemistry and Molecular Biology (1980)				
	AT	6	Martin, C.T., et al., "Processivity in Early Stages of Transcription by T7 RNA Polymerase," Biochemistry 27:3966-3974, American Chemical Society (1988)				

EXAMINER 	DATE CONSIDERED 12-18-06
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	AL						Yes No
	AM						Yes No
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	2	Meyer, K.L., and Hanna, M.M., "Synthesis and Characterization of a New 5-Thiol-Protected Deoxyuridine Phosphoramidite for Site-Specific Modification of DNA," Bioconjugate Chem. 7:401-412, American Chemical Society (1996)
	AS	2	Milligan, J.F., and Uhlenbeck, O.C., "Synthesis of Small RNAs Using T7 RNA Polymerase," Methods Enzymol. 180:51-62, Academic Press, Inc. (1989)
	AT	2	Montemagno, C., and Bachand, G., "Constructing nanomechanical devices powered by biomolecular motors," Nanotechnology 10:225-231, IOP Publishing, Ltd. (1999)

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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	8	Mullis, K., et al., "Specific Enzymatic Amplification of DNA In Vitro: The Polymerase Chain Reaction," Cold Spring Harb. Symp. Quant. Biol. 51:263-273, Cold Spring Harbor Laboratory (1986)
	AS	8	Mullis, K.B., et al., "The Polymerase Chain Reaction: Why It Works," in Polymerase Chain Reaction, Erlich, H.A., et al., eds., Cold Spring Harbor Laboratory Press, Woodbury, N.Y., pp. 237-243 (1989)
	AT	8	Nelson, P.S., et al., "Bifunctional oligonucleotide probes synthesized using a novel CPG support are able to detect single base pair mutations," Nucleic Acids Res. 17:7187-7194, Oxford University Press (1989)

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	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	2	Pringsheim, P., "Fluorescence of organic compounds," in Fluorescence and phosphorescence, Pringsham, P., ed., Interscience Publishers, Inc. New York, N.Y., pp. 392-397, 420-423 (1949)
	AS	2	Saiki, R.K., et al., "Primer-Directed Enzymatic Amplification of DNA with a Thermostable DNA Polymerase," Science 239:487-491, American Association for the Advancement of Science (1988)
	AT	2	Sinha, N.D., and Striepeke, S., "Oligonucleotides with reporter groups attached to the 5'-terminus," in Oligonucleotides and Analogues: A Practical Approach, Eckstein, F., ed., Oxford University Press pp. 185-189, 200-201 (1992)

EXAMINER

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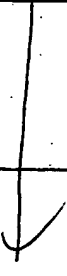

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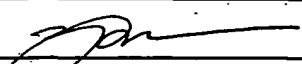
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
	AR	<u>10</u>	Smithies, O., et al., "Detection of Targeted Gene Modifications by Polymerase Chain Reaction," in Polymerase Chain Reaction, Erlich, H.A., et al., eds., Cold Spring Harbor Laboratory Press, Woodbury, N.Y., pp. 199-203 (1989)
	AS	<u>10</u>	Sproat, B.S., et al., "The synthesis of protected 5'-mercapto-2',5'-dideoxyribonucleoside-3'-O-phosphoramidites; uses of 5'-mercapto-oligodeoxyribonucleotides," Nucleic Acids Res. 15:4837-4848, Oxford University Press (1987)
	AT	<u>10</u>	Walker, G.T., et al., "Isothermal in vitro amplification of DNA by a restriction enzyme/DNA polymerase system," Proc. Natl. Acad. Sci. USA 89:392-396, National Academy Press (1992)

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	AR	11	Zuckermann, R., et al., "Efficient methods for attachment of thiol specific probes to the 3'-ends of synthetic oligodeoxyribonucleotides," Nucleic Acids Res. 15:5305-5321, Oxford University Press (1987)
	AS		Daube et al., "Functional Transcription Elongation Complexes from Synthetic RNA-DNA Bubble Complexes," Science, 1992, vol. 258, pages 1320-1324
	AT		Mancini et al., "Constitutively Methylated CpG Dinucleotides as Mutation Hot Spots in the Retinoblastoma Gene (RB1), American Journal of Human Genetics, 1997, vol. 61, pages 80-87

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10/602,045APPLICANT
Michelle M. HannaFILING DATE
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	AA3	5,399,491	03/21/95	Kacian <i>et al.</i>			03/19/92
	AB3	5,888,819	03/30/99	Goelet <i>et al.</i>			03/05/91
	AC3	5,554,516	09/10/96	Kacian <i>et al.</i>			12/02/93
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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	12	International Search Report for International Application No. PCT/US02/34419, mailed July 24, 2003, International Searching Authority/USA, Washington, D.C.
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	AA						
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	AD3	5,571,669	11/05/1996	Lu et al.			01/14/1993
	AE3	6,251,594 B1	06/26/2001	Gonzalgo et al.			06/09/1998
	AF3	2003/0099950 A1	05/29/2003	Hanna, M.M.			10/30/2001
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FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
	AL1	WO 89/01050	02/09/1989	WIPO			Yes No
	AM1	WO 03/038042 A2	05/08/2003	WIPO			Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR		
	AS	<u>12</u>	Antequerra, F., and Bird, A., "Number of CpG islands and genes in human and mouse," Proc. Natl. Acad. Sci. USA 90:11995-11999, National Academy of Sciences (1993).
	AT	<u>12</u>	Callahan, R., et al., "Frequent Mutations in Breast Cancer," Ann. N.Y. Acad. Sci. 698:21-30, New York Academy of Sciences (1993).

EXAMINER

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EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to Applicant.



FORM PTO-1449 <u>SECOND SUPPLEMENTAL</u> <u>INFORMATION DISCLOSURE STATEMENT</u>	ATTY. DOCKET NO. 2072.0010003/LBB/SJE	APPLICATION NO. 10/602,045
	APPLICANT Michelle Hanna	
	FILING DATE June 24, 2003	GROUP To be assigned

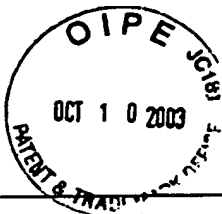
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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
	AR	13	Cifone, M.A., and Fidler, I.J., "Increasing metastatic potential is associated with increasing genetic instability of clones isolated from murine neoplasms," Proc. Natl. Acad. Sci. USA 78:6949-6952, National Academy of Sciences (1981).
	AS	13	Esteller, M., et al., "Promoter Hypermethylation and BRCA1 Inactivation in Sporadic Breast and Ovarian Tumors," J. Natl. Cancer Inst. 92:564-569, Oxford University Press (April 2000). (Retrieved from LexisNexis Academic database)
	AT	13	Frommer, M., et al., "A genomic sequencing protocol that yields a positive display of 5-methylcytosine residues in individual DNA strands," Proc. Natl. Acad. Sci. USA 89:1827-1831, National Academy of Sciences (1992).

EXAMINER 	DATE CONSIDERED 2/16/05
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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)				
	AR	<u>14</u>	Giusti, W.G., and Adriano, T., "Synthesis and Characterization of 5'-Fluorescent-dye-labeled Oligonucleotides," <i>PCR Methods Appl.</i> 2:223-227, Cold Spring Harbor Laboratory Press (1993).	
	AS	<u>14</u>	Gupta, K.C., et al., "A general method for the synthesis of 3'-sulfhydryl and phosphate group containing oligonucleotides," <i>Nucleic Acids Res.</i> 19:3019-3025, Oxford University Press (1991).	
	AT	<u>14</u>	Herman, J.G., et al., "Inactivation of the <i>CDKN2/p16/MTS1</i> Gene Is Frequently Associated with Aberrant DNA Methylation in All Common Human Cancers," <i>Cancer Res.</i> 55:4525-4530, American Association for Cancer Research (1995).	

EXAMINER 	DATE CONSIDERED 2/16/06
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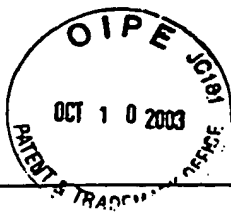
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>15</u>	Herman, J.G., et al., "Incidence and functional consequences of hMLH1 promoter hypermethylation in colorectal carcinoma," Proc. Natl. Acad. Sci. USA 95:6870-6875, National Academy of Sciences (1998).
	AS	<u>15</u>	Horii, A., et al., "Frequent Replication Errors at Microsatellite Loci in Tumors of Patients with Multiple Primary Cancers," Cancer Res. 54:3373-3375, American Association for Cancer Research (1994).
	AT	<u>15</u>	Issa, J.-P.J., et al., "Methylation of the oestrogen receptor CpG island links ageing and neoplasia in human colon," Nat. Genet. 7:536-540, Nature Publishing Co. (1994).

EXAMINER 	DATE CONSIDERED 2-16-06
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FORM PTO-1449 SECOND SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT	ATTY. DOCKET NO. 2072.0010003/LBE/SJE	APPLICATION NO. 10/602,045
	APPLICANT Michelle Hanna	
	FILING DATE June 24, 2003	GROUP To be assigned

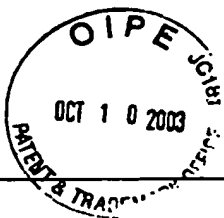
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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	16	Jin, D.J., "An Escherichia coli RNA Polymerase Defective in Transcription due to its Overproduction of Abortive Initiation Products," J. Mol. Biol. 236:72-80, Academic Press Limited (1994).
	AS	16	Langer, P.R., et al., "Enzymatic synthesis of biotin-labeled polynucleotides: Novel nucleic acid affinity probes," Proc. Natl. Acad. Sci. USA 78:6633-6637, National Academy of Sciences (1981).
	AT	16	Loeb, L.A., "Mutator Phenotype May Be Required for Multistage Carcinogenesis," Cancer Res. 51:3075-3079, American Association for Cancer Research (1991).

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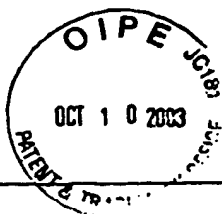
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	AO						Yes No
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	17	Meinkoth, J., and Wahl, G., "Hybridization of Nucleic Acids Immobilized on Solid Supports," <i>Anal. Biochem.</i> 138:267-284, Academic Press, Inc. (1984).
	AS	17	Nowell, P.C., "The Clonal Evolution of Tumor Cell Populations," <i>Science</i> 194:23-28, American Association for the Advancement of Science (1976).
	AT	17	Palmisano, W.A., et al., "Predicting Lung Cancer by Detecting Aberrant Promoter Methylation in Sputum," <i>Cancer Res.</i> 60:5954-5958, American Association for Cancer Research (November 2000).

EXAMINER 	DATE CONSIDERED 2-16-06
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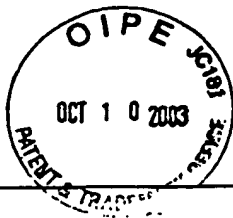
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>18</u>	Radlowski, M., and Job, D., "Effect of disulfide and sulfhydryl reagents on abortive and productive elongation catalyzed by <i>Escherichia coli</i> RNA polymerase," <i>Acta Biochim. Pol.</i> 41:415-419, Panstwowe Wydawnictwo Naukowe (1994).
	AS	<u>18</u>	Rice, J.C., et al., "Aberrant methylation of the <i>BRCA1</i> CpG island promoter is associated with decreased <i>BRCA1</i> mRNA in sporadic breast cancer cells," <i>Oncogene</i> 17:1807-1812, Stockton Press (1998).
	AT	<u>18</u>	Robertson, K.D., "DNA methylation, methyltransferases, and cancer," <i>Oncogene</i> 20:3139-3155, Nature Publishing Group (May 2001).

EXAMINER 	DATE CONSIDERED 2-1-06
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	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	19	Tlsty, T.D., et al., "Differences in the rates of gene amplification in nontumorigenic and tumorigenic cell lines as measured by Luria-Delbrück fluctuation analysis," <i>Proc. Natl. Acad. Sci. USA</i> 86:9441-9445, National Academy of Sciences (1989).
	AS	19	Toyota, M., and Issa, J.-P.J., "CpG island methylator phenotypes in aging and cancer," <i>Semin. Cancer Biol.</i> 9:349-357, Academic Press (1999).
	AT	19	Toyota, M., and Issa, J.-P.J., "The role of DNA hypermethylation in human neoplasia," <i>Electrophoresis</i> 21:329-333, Wiley-VCH Verlag GmbH (January 2000).

EXAMINER 	DATE CONSIDERED 2-11-06
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	20	Vogelstein, B., and Kinzler, K.W., "The multistep nature of cancer," Trends Genet. 9:138-141, Elsevier Science Publishers Ltd. (1993).
	AS	20	Wang, J.-Y., et al., "Monovalent cations differ in their effects on transcription initiation from a σ -70 promoter of <i>Escherichia coli</i> ," Gene 196:95-98, Elsevier Science B.V. (1997).
	AT	20	Wu, D.Y., and Wallace, R.B., "The Ligation Amplification Reaction (LAR) - Amplification of Specific DNA Sequences Using Sequential Rounds of Template-Dependent Ligation," Genomics 4:560-569, Academic Press, Inc. (1989).

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